

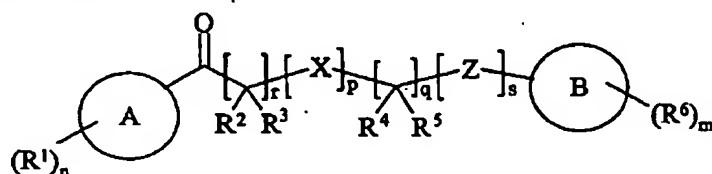
Application No.: Not Yet Assigned

Docket No.: ASZD-P01-804

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

1. (Currently Amended) A method for inhibiting 11 $\beta$ HSD1, comprising administering a compound of formula (I):



(I)

wherein:

**Ring A** is selected from aryl or heteroaryl;

**R¹** is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, C<sub>1-6</sub>alkyl, C<sub>2-6</sub>alkenyl, C<sub>2-6</sub>alkynyl, C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>alkanoyl, C<sub>1-6</sub>alkanoyloxy, *N*-(C<sub>1-6</sub>alkyl)amino, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkanoylamino, *N*-(C<sub>1-6</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-6</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-6</sub>alkoxycarbonyl, *N*-(C<sub>1-6</sub>alkyl)sulphamoyl, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>sulphamoyl, C<sub>1-6</sub>alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclylC<sub>0-6</sub>alkylene-Y-, and heterocyclylC<sub>0-6</sub>alkylene-Y-; or two **R¹** groups on adjacent carbons may form an oxyC<sub>1-4</sub>alkoxy group or a C<sub>3-5</sub>alkylene group; wherein **R¹** may be optionally substituted on carbon by-with one or more **R⁷** groups-selected ~~from-R⁷~~; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by-with an **R⁸** group-selected ~~from-R⁸~~;

**n** is 0-3; wherein the values of **R¹** may be the same or different;

**R², R³, R⁴, and R⁵** are independently selected from hydrogen, hydroxy, amino, cyano, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkoxy, *N*-(C<sub>1-4</sub>alkyl)amino, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-4</sub>alkoxycarbonyl, C<sub>1-4</sub>alkoxycarbonylamino, C<sub>1-4</sub>alkanoyloxy, carbocyclyl, heterocyclyl, carbocyclylC<sub>1-4</sub>alkyl, and heterocyclylC<sub>1-4</sub>alkyl; or **R²** and **R³** together form oxo or a spiro attached heterocyclyl; wherein **R², R³, R⁴, and R⁵** may be independently optionally substituted on carbon by-with one or more **R⁹** groups-selected ~~from-R⁹~~; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by-with an **R¹⁰** group-selected ~~from-R¹⁰~~;

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X and Z are independently selected from  $-CR^{11}R^{12}-$ ,  $-S(O)_a-$ ,  $-O-$ ,  $-NR^{13}-$ ,  $-C(O)-$ ,  $-C(O)NR^{14}-$ ,  $-NR^{15}C(O)-$ ,  $-OC(O)-$ ,  $-C(O)O-$ ,  $-SO_2NR^{16}-$ , ~~or~~ and  $-NR^{16}SO_2-$ ; wherein a is 0 to 2;

r is 1 or 2;

q is 0 or 1;

p is 0 or 1;

s is 0 or 1;

Ring B is carbocyclyl or heterocyclyl; wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by an  $R^{17}$  group ~~selected from  $R^{17}$~~ ;

$R^6$  is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy,  $C_{1-4}$ alkyl,  $C_{2-4}$ alkenyl,  $C_{2-4}$ alkynyl,  $C_{1-4}$ alkoxy,  $C_{1-4}$ alkanoyl,  $C_{1-4}$ alkanoyloxy,  $N-(C_{1-4}$ alkyl)amino,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkanoylamino,  $N-(C_{1-4}$ alkyl)carbamoyl,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-4}$ alkylS(O)<sub>a</sub> wherein a is 0 to 2,  $C_{1-4}$ alkoxycarbonyl,  $N-(C_{1-4}$ alkyl)sulphamoyl,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>sulphamoyl,  $C_{1-4}$ alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclylC<sub>0-4</sub>alkylene-Y-, and heterocyclylC<sub>0-4</sub>alkylene-Y-; wherein  $R^6$  may be optionally substituted on carbon by with one or more  $R^{18}$  groups ~~selected from  $R^{18}$~~ ; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by with an  $R^{19}$  group ~~selected from  $R^{19}$~~ ;

m is 0-3; wherein the values of  $R^6$  may be the same or different;

Y is  $-S(O)_a-$ ,  $-O-$ ,  $-NR^{20}-$ ,  $-C(O)-$ ,  $-C(O)NR^{21}-$ ,  $-NR^{22}C(O)-$ , or  $-SO_2NR^{23}-$ ; wherein a is 0 to 2;

$R^7$ ,  $R^9$ , and  $R^{18}$  are independently selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy,  $C_{1-4}$ alkyl,  $C_{2-4}$ alkenyl,  $C_{2-4}$ alkynyl,  $C_{1-4}$ alkoxy,  $C_{1-4}$ alkanoyl,  $C_{1-4}$ alkanoyloxy,  $N-(C_{1-4}$ alkyl)amino,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkanoylamino,  $N-(C_{1-4}$ alkyl)carbamoyl,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-4}$ alkylS(O)<sub>a</sub> wherein a is 0 to 2,  $C_{1-4}$ alkoxycarbonyl,  $N-(C_{1-4}$ alkyl)sulphamoyl,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>sulphamoyl,  $C_{1-4}$ alkylsulphonylamino, carbocyclyl, and heterocyclyl; wherein  $R^7$ ,  $R^9$ , and  $R^{18}$  may be independently optionally substituted on carbon by with one or more  $R^{26}$  groups;

$R^{11}$  and  $R^{12}$  are independently selected from hydrogen, hydroxy, amino, cyano,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $N-(C_{1-4}$ alkyl)amino,  $N,N-(C_{1-4}$ alkyl)<sub>2</sub>amino, carbocyclyl, heterocyclyl, carbocyclylC<sub>1-4</sub>alkyl, and heterocyclylC<sub>1-4</sub>alkyl; wherein  $R^{11}$  and  $R^{12}$  may be independently

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optionally substituted on carbon ~~by with~~ one or more R<sup>24</sup> groups selected from R<sup>24</sup>; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by with~~ an R<sup>25</sup> group selected from R<sup>25</sup>;

R<sup>24</sup> is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy, C<sub>1-4</sub>alkyl, C<sub>2-4</sub>alkenyl, C<sub>2-4</sub>alkynyl, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkanoyloxy, *N*-(C<sub>1-4</sub>alkyl)amino, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkanoylamino, *N*-(C<sub>1-4</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-4</sub>alkoxycarbonyl, *N*-(C<sub>1-4</sub>alkyl)sulphamoyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>sulphamoyl, and C<sub>1-4</sub>alkylsulphonylamino;

R<sup>8</sup>, R<sup>10</sup>, R<sup>17</sup>, R<sup>19</sup>, and R<sup>25</sup> are independently selected from C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkylsulphonyl, C<sub>1-4</sub>alkoxycarbonyl, carbamoyl, *N*-(C<sub>1-4</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, benzyl, benzyloxycarbonyl, benzoyl, carbocyclyl, heterocyclyl, and phenylsulphonyl; wherein R<sup>8</sup>, R<sup>10</sup>, R<sup>17</sup>, R<sup>19</sup>, and R<sup>25</sup> may be independently optionally substituted on carbon ~~by with~~ one or more R<sup>27</sup> groups;

R<sup>13</sup>, R<sup>14</sup>, R<sup>15</sup>, R<sup>16</sup>, R<sup>20</sup>, R<sup>21</sup>, R<sup>22</sup>, and R<sup>23</sup> are independently selected from hydrogen, phenyl, C<sub>1-4</sub>alkylsulphonyl, and C<sub>1-4</sub>alkyl;

R<sup>26</sup> and R<sup>27</sup> are independently selected from selected from halo, nitro, cyano, hydroxy, trifluoromethoxy, trifluoromethyl, amino, carboxy, carbamoyl, mercapto, sulphamoyl, methyl, ethyl, methoxy, ethoxy, acetyl, acetoxyl, methylamino, ethylamino, dimethylamino, diethylamino, *N*-methyl-*N*-ethylamino, acetylaminyl, *N*-methylcarbamoyl, *N*-ethylcarbamoyl, *N,N*-dimethylcarbamoyl, *N,N*-diethylcarbamoyl, *N*-methyl-*N*-ethylcarbamoyl, methylthio, ethylthio, methylsulphanyl, ethylsulphanyl, mesyl, ethylsulphonyl, methoxycarbonyl, ethoxycarbonyl, *N*-methylsulphamoyl, *N*-ethylsulphamoyl, *N,N*-dimethylsulphamoyl, *N,N*-diethylsulphamoyl, and or *N*-methyl-*N*-ethylsulphamoyl;

or a pharmaceutically acceptable salt thereof;

~~in the manufacture of a medicament for use in the inhibition of 11βHSD1;~~

with the proviso that said compound is not (1-methyl-1-pyrid-3-ylethyl)-(pyrid-3-yl)-ketone.

2. (Currently Amended) The ~~method use of a compound, or a pharmaceutically acceptable salt thereof, as claimed in of~~ claim 1, wherein Ring A is selected from phenyl, naphthyl, thienyl, furyl, thiazolyl, pyridyl, imidazolyl, benzothiazolyl, and or benzothienyl.

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3. (Currently Amended) The ~~methoduse of a compound, or a pharmaceutically acceptable salt thereof, as claimed in either of claim 1, or claim 2~~ wherein  $R^1$  is selected from halo, cyano, hydroxy,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkylsulphonylamino, carbocyclyl, and heterocyclyl $C_{0-4}$ alkylene-Y-; or two  $R^1$  groups on adjacent carbons may form an oxy $C_{1-4}$ alkoxy group; wherein  $R^1$  may be optionally substituted on carbon by with one or more  $R^7$  groups selected from  $R^7$ ;

Y is -S(O)<sub>a</sub>-, or -O-; wherein a is 0 to 2; and

$R^7$  is halo.

4. (Currently Amended) The ~~methoduse of a compound, or a pharmaceutically acceptable salt thereof, as claimed in any one of claims 1, [[-3]]~~ wherein  $R^2$ ,  $R^3$ ,  $R^4$ , and  $R^5$  are independently selected from hydrogen, hydroxy,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $N$ -( $C_{1-4}$ alkyl)amino, carbocyclyl, carbocyclyl $C_{1-4}$ alkyl, and heterocyclyl $C_{1-4}$ alkyl; wherein  $R^2$ ,  $R^3$ ,  $R^4$ , and  $R^5$  may be independently optionally substituted on carbon by with one or more  $R^9$  groups selected from  $R^9$ ; and wherein

$R^9$  is selected from halo, cyano,  $C_{1-4}$ alkyl, and  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino.

5. (Currently Amended) The ~~methoduse of a compound, or a pharmaceutically acceptable salt thereof, as claimed in any one of claims 1, [[-6]]~~ wherein X is -S(O)<sub>a</sub>-, -O-, -NR<sup>13</sup>-, -NR<sup>15</sup>C(O)-, -SO<sub>2</sub>NR<sup>16</sup>-, or -NR<sup>16</sup>SO<sub>2</sub>-; wherein a is 0 or 2; and

$R^{13}$ ,  $R^{15}$ , and  $R^{16}$  are independently selected from hydrogen, phenyl,  $C_{1-4}$ alkylsulphonyl, and  $C_{1-4}$ alkyl.

6. (Currently Amended) The ~~methoduse of a compound, or a pharmaceutically acceptable salt thereof, as claimed in any one of claims 1, [[-5]]~~ wherein Ring B is phenyl, thienyl, furyl, thiazolyl, piperidinyl, piperazinyl, pyrrolidinyl, 1,3-dihydroisoindolyl, morpholinyl, naphthyl, cyclohexyl, pyridyl, imidazolyl, 1,2,4-triazolyl, 1,3-benzodioxolyl, thiomorpholinyl, pyrimidinyl, pyrazinyl, pyridazinyl, benzimidazolyl, or pyrimidinyl; wherein if Ring B contains an -NH- moiety, that nitrogen may be optionally substituted by with an  $R^{17}$  group selected from  $R^{17}$ ;

$R^{17}$  is  $C_{1-4}$ alkyl or benzyl; wherein  $R^{17}$  may be optionally substituted on carbon by with one or more  $R^{27}$  groups; wherein and

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R<sup>27</sup> is methoxy.

7. (Currently Amended) The ~~methoduse of a compound, or a pharmaceutically acceptable salt thereof, as claimed in any one of claims 1, [[-6]]~~ wherein R<sup>6</sup> is a substituent on carbon and is selected from halo, hydroxy, nitro, cyano, carbamoyl, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkanoyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkanoylamino, *N*-(C<sub>1-4</sub>alkyl)carbamoyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 or 2, C<sub>1-4</sub>alkoxycarbonyl, *N,N*-(C<sub>1-4</sub>alkyl)<sub>2</sub>sulphamoyl, carbocyclyl, heterocyclyl, and carbocyclylC<sub>0-4</sub>alkylene-Y-; wherein R<sup>6</sup> may be optionally substituted on carbon ~~by with~~ one or more R<sup>18</sup> groups ~~selected from R<sup>18</sup>~~; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by with an R<sup>19</sup> group selected from R<sup>19</sup>~~;

Y is -C(O) or -C(O)NR<sup>21</sup>-;

R<sup>18</sup> is selected from halo, cyano, hydroxy, C<sub>1-4</sub>alkoxy, and heterocyclyl;

R<sup>19</sup> is heterocyclyl; and

R<sup>21</sup> is hydrogen.

8. (Currently Amended) The ~~methoduse of a compound of formula (I) (as depicted in claim 1, [D])~~ wherein:

Ring A is selected from phenyl, naphthyl, thienyl, furyl, thiazolyl, pyridyl, imidazolyl, benzothiazolyl, and ~~or~~ benzothienyl;

R<sup>1</sup> is selected from halo, cyano, hydroxy, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, *N,N*-(C<sub>1-6</sub>alkyl)<sub>2</sub>amino, C<sub>1-6</sub>alkylsulphonylamino, carbocyclyl, and heterocyclylC<sub>0-6</sub>alkylene-Y-; or two R<sup>1</sup> groups on adjacent carbons may form an oxyC<sub>1-4</sub>alkoxy group; wherein R<sup>1</sup> may be optionally substituted on carbon ~~by with~~ one or more R<sup>7</sup> groups ~~selected from R<sup>7</sup>~~;

Y is -S(O)<sub>a</sub>-, or -O-; wherein a is 0 to 2; and

R<sup>7</sup> is halo[.];

n is 0-3; wherein the values of R<sup>1</sup> may be the same or different;

r is 1 or 2;

s is 0;

R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, and R<sup>5</sup> are independently selected from hydrogen, hydroxy, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkoxy, *N*-(C<sub>1-4</sub>alkyl)amino, carbocyclyl, carbocyclylC<sub>1-4</sub>alkyl, and heterocyclylC<sub>1-4</sub>alkyl;

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wherein  $R^2$ ,  $R^3$ ,  $R^4$ , and  $R^5$  may be independently optionally substituted on carbon ~~by with~~ one or more  $R^9$  ~~groups selected from  $R^9$~~ ; wherein

$R^9$  is selected from halo, cyano,  $C_{1-4}$ alkyl, and  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino[.];

$X$  is  $-S(O)_a$ -,  $-O$ -,  $-NR^{13}$ -,  $-NR^{15}C(O)$ -,  $-SO_2NR^{16}$ -, or  $-NR^{16}SO_2$ -; wherein  $a$  is 0 or 2; and

$R^{13}$ ,  $R^{15}$ , and  $R^{16}$  are independently selected from hydrogen, phenyl,  $C_{1-4}$ alkylsulphonyl, and  $C_{1-4}$ alkyl;

$q$  is 0 or 1;

$p$  is 0 or 1;

Ring B is phenyl, thienyl, furyl, thiazolyl, piperidinyl, piperazinyl, pyrrolidinyl, 1,3-dihydroisindolyl, morpholinyl, naphthyl, cyclohexyl, pyridyl, imidazolyl, 1,2,4-triazolyl, 1,3-benzodioxolyl, thiomorpholinyl, pyrimidinyl, pyrazinyl, pyridazinyl, benzimidazolyl, or pyrimidinyl; wherein if Ring B contains an  $-NH$ - moiety, that nitrogen may be optionally substituted by a group selected from  $R^{17}$ ;

$R^{17}$  is  $C_{1-4}$ alkyl or benzyl; wherein  $R^{17}$  may be optionally substituted on carbon ~~by with~~ one or more  $R^{27}$  ~~groups; wherein~~

$R^{27}$  is methoxy;

$R^6$  is a substituent on carbon and is selected from halo, hydroxy, nitro, cyano, carbamoyl,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $C_{1-4}$ alkanoyl,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkanoylamino,  $N$ -( $C_{1-4}$ alkyl)carbamoyl,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-4}$ alkyl $S(O)_a$  wherein  $a$  is 0 or 2,  $C_{1-4}$ alkoxycarbonyl,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>sulphamoyl, carbocyclyl, heterocyclyl, and carbocyclyl $C_{0-4}$ alkylene- $Y$ -; wherein  $R^6$  may be optionally substituted on carbon ~~by with~~ one or more  $R^{18}$  ~~groups selected from  $R^{18}$~~ ; and wherein if said heterocyclyl contains an  $-NH$ - moiety, that nitrogen may be optionally substituted ~~by with~~ an  $R^{19}$  ~~group selected from  $R^{19}$~~ ;

$Y$  is  $-C(O)$  or  $-C(O)NR^{21}$ ;

$R^{18}$  is selected from halo, cyano, hydroxy,  $C_{1-4}$ alkoxy, and heterocyclyl;

$R^{19}$  is heterocyclyl; ~~and~~

$R^{21}$  is hydrogen; ~~and~~

$m$  is 0-3; wherein the values of  $R^6$  may be the same or different[.];]

~~or a pharmaceutically acceptable salt thereof;~~

~~in the manufacture of a medicament for use in the inhibition of 11 $\beta$ HSD1;~~

~~with the proviso that said compound is not (1-methyl-1-pyrid-3-ylethyl)-(pyrid-3-yl)-ketone.~~

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9. (Currently Amended) A compound of formula (I) (as depicted in claim 1) selected from:

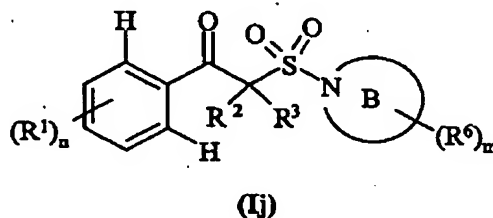
[2-(4-chlorophenyl)-1-(pyrid-3-yl)ethyl]-(4-chlorophenyl)-ketone;  
 [2-(4-chlorophenyl)-1-(pyrazin-2-yl)ethyl]-(pyridin-3-yl)-ketone;  
 ( $\alpha$ -methylamino-4-chlorobenzyl)-(4-chlorophenyl)-ketone;  
 (benzothiazol-2-yl)-(pyrrolidin-1-ylsulphonylmethyl)-ketone;  
 (thiazol-2-yl)-(pyrrolidin-1-ylsulphonylmethyl)-ketone;  
 [1-(morpholinosulphonyl)-1-methylethyl]-(4-fluorophenyl)-ketone;  
 (4-fluorophenyl)-[*N*-(cyclohexyl)-*N*-(isopropyl)sulphamoylmethyl]-ketone;  
 (4-fluorophenyl)-[*N*-(pyrid-2-yl)-*N*-(methyl)sulphamoylmethyl]-ketone;  
 (4-methylphenylsulphonylmethyl)-(4-cyanophenyl)-ketone;  
 (4-ethoxyphenoxyethyl)-(4-chlorophenyl)-ketone;  
 (4-chlorophenyl)-[3-(2,6-difluorobenzoylamino) propyl]-ketone; and  
 (4-chlorophenyl)-[3-(4-methoxyphenylsulphonylamino)propyl]-ketone;  
 or a pharmaceutically acceptable salt thereof.

10. (Currently Amended) The ~~methoduse of a compound of formula (I) (as depicted in claim 1, [I])~~ wherein the compound of formula (I) is selected from:

( $\alpha$ -methyl- $\alpha$ -hydroxy-4-chlorobenzyl)-(4-chlorophenyl)-ketone;  
 (morpholinosulphonylmethyl)-(4-fluorophenyl)-ketone;  
 (*N*-methyl-4-methylanilinosulphonylmethyl)-(4-chlorophenyl)-ketone; and  
 (*N*-methyl-4-chloroanilinomethyl)-(4-chlorophenyl)-ketone;  
 or a pharmaceutically acceptable salt thereof[[:]]

~~in the manufacture of a medicament for use in the inhibition of 11 $\beta$ HSD1.~~

11. (Currently Amended) A compound of formula (Ij):



wherein:

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$R^1$  is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl,  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{1-6}$ alkoxy,  $C_{1-6}$ alkanoyl,  $C_{1-6}$ alkanoyloxy,  $N$ -( $C_{1-6}$ alkyl)amino,  $N,N$ -( $C_{1-6}$ alkyl)<sub>2</sub>amino,  $C_{1-6}$ alkanoylamino,  $N$ -( $C_{1-6}$ alkyl)carbamoyl,  $N,N$ -( $C_{1-6}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-6}$ alkylS(O)<sub>*a*</sub> wherein *a* is 0 to 2,  $C_{1-6}$ alkoxycarbonyl,  $N$ -( $C_{1-6}$ alkyl)sulphamoyl,  $N,N$ -( $C_{1-6}$ alkyl)<sub>2</sub>sulphamoyl,  $C_{1-6}$ alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclylC<sub>0-6</sub>alkylene-Y-, and heterocyclylC<sub>0-6</sub>alkylene-Y-; or two  $R^1$  groups on adjacent carbons may form an oxyC<sub>1-4</sub>alkoxy group or a C<sub>3-5</sub>alkylene group; wherein  $R^1$  may be optionally substituted on carbon by with one or more  $R^7$  groups ~~selected from  $R^7$~~ ; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by an  $R^8$  group ~~selected from  $R^8$~~ ;

*n* is 0-3; wherein the values of  $R^1$  may be the same or different;

$R^2$  and  $R^3$  are independently selected from hydrogen, hydroxy, amino, cyano,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $N$ -( $C_{1-4}$ alkyl)amino,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkylS(O)<sub>*a*</sub> wherein *a* is 0 to 2,  $C_{1-4}$ alkoxycarbonyl,  $C_{1-4}$ alkoxycarbonylamino,  $C_{1-4}$ alkanoyloxy, carbocyclyl, heterocyclyl, carbocyclylC<sub>1-4</sub>alkyl, and heterocyclylC<sub>1-4</sub>alkyl; or

$R^2$  and  $R^3$  together form oxo or a spiro attached heterocyclyl; wherein  $R^2$  and  $R^3$  may be independently optionally substituted on carbon by with one or more  $R^9$  groups ~~selected from  $R^9$~~ ; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by with an  $R^{10}$  group ~~selected from  $R^{10}$~~ ;

Ring B is a heterocyclyl linked to the sulphonyl of the compound of formula (Ij) via a nitrogen atom; wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by with an  $R^{17}$  group ~~selected from  $R^{17}$~~ ;

$R^6$  is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy,  $C_{1-4}$ alkyl,  $C_{2-4}$ alkenyl,  $C_{2-4}$ alkynyl,  $C_{1-4}$ alkoxy,  $C_{1-4}$ alkanoyl,  $C_{1-4}$ alkanoyloxy,  $N$ -( $C_{1-4}$ alkyl)amino,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>amino,  $C_{1-4}$ alkanoylamino,  $N$ -( $C_{1-4}$ alkyl)carbamoyl,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>carbamoyl,  $C_{1-4}$ alkylS(O)<sub>*a*</sub> wherein *a* is 0 to 2,  $C_{1-4}$ alkoxycarbonyl,  $N$ -( $C_{1-4}$ alkyl)sulphamoyl,  $N,N$ -( $C_{1-4}$ alkyl)<sub>2</sub>sulphamoyl,  $C_{1-4}$ alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclylC<sub>0-4</sub>alkylene-Y-, and heterocyclylC<sub>0-4</sub>alkylene-Y-; wherein  $R^6$  may be optionally substituted on carbon by with one or more  $R^{18}$  groups ~~selected from  $R^{18}$~~ ; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted by with an  $R^{19}$  group ~~selected from  $R^{19}$~~ ;



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m is 0-3; wherein the values of R<sup>6</sup> may be the same or different;

Y is -S(O)<sub>a</sub>-, -O-, -NR<sup>20</sup>-, -C(O)-, -C(O)NR<sup>21</sup>-, -NR<sup>22</sup>C(O)-, or -SO<sub>2</sub>NR<sup>23</sup>-; wherein a is 0 to 2;

R<sup>7</sup>, R<sup>9</sup>, and R<sup>18</sup> are independently selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy, C<sub>1-4</sub>alkyl, C<sub>2-4</sub>alkenyl, C<sub>2-4</sub>alkynyl, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkanoyloxy, N-(C<sub>1-4</sub>alkyl)amino, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkanoylamino, N-(C<sub>1-4</sub>alkyl)carbamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-4</sub>alkoxycarbonyl, N-(C<sub>1-4</sub>alkyl)sulphamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>sulphamoyl, C<sub>1-4</sub>alkylsulphonylamino, carbocyclyl, and heterocyclyl; wherein R<sup>7</sup>, R<sup>9</sup>, and R<sup>18</sup> may be independently optionally substituted on carbon ~~by~~with one or more R<sup>26</sup> groups;

R<sup>8</sup>, R<sup>10</sup>, R<sup>17</sup>, and R<sup>19</sup> are independently selected from C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkylsulphonyl, C<sub>1-4</sub>alkoxycarbonyl, carbamoyl, N-(C<sub>1-4</sub>alkyl)carbamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, benzyl, benzyloxycarbonyl, benzoyl, carbocyclyl, heterocyclyl, and phenylsulphonyl; wherein R<sup>8</sup>, R<sup>10</sup>, R<sup>17</sup>, and R<sup>19</sup> may be independently optionally substituted on carbon ~~by~~with one or more R<sup>27</sup> groups;

R<sup>20</sup>, R<sup>21</sup>, R<sup>22</sup>, and R<sup>23</sup> are independently selected from hydrogen, phenyl, C<sub>1-4</sub>alkylsulphonyl, and C<sub>1-4</sub>alkyl;

R<sup>26</sup> and R<sup>27</sup> are independently selected from selected from halo, nitro, cyano, hydroxy, trifluoromethoxy, trifluoromethyl, amino, carboxy, carbamoyl, mercapto, sulphamoyl, methyl, ethyl, methoxy, ethoxy, acetyl, acetoxyl, methylamino, ethylamino, dimethylamino, diethylamino, N-methyl-N-ethylamino, acetylamino, N-methylcarbamoyl, N-ethylcarbamoyl, N,N-dimethylcarbamoyl, N,N-diethylcarbamoyl, N-methyl-N-ethylcarbamoyl, methylthio, ethylthio, methylsulphinyl, ethylsulphinyl, mesyl, ethylsulphonyl, methoxycarbonyl, ethoxycarbonyl, N-methylsulphamoyl, N-ethylsulphamoyl, N,N-dimethylsulphamoyl, N,N-diethylsulphamoyl, and ~~or~~ N-methyl-N-ethylsulphamoyl;

or a pharmaceutically acceptable salt thereof;

with the proviso that said compound is not

(phenyl)-[α-(pyrrolidin-1-ylsulphonyl)benzyl]-ketone;

(phenyl)-[α-(morpholinosulphonyl)benzyl]-ketone;

(4-carbamoylphenyl)-[4-(5-chloropyridin-2-yloxy)piperidin-1-ylsulphonylmethyl]-ketone;

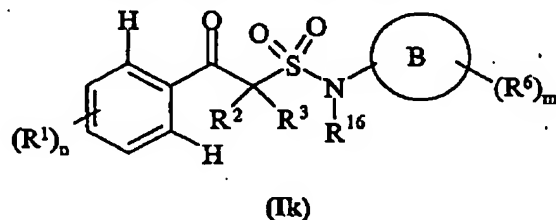
(4-carbamoylphenyl)-[4-(4-fluorophenyl)piperidin-1-ylsulphonylmethyl]-ketone;

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(4-fluorophenyl)-[4-(5-chloropyridin-2-yloxy)piperidin-1-ylsulphonylmethyl]-ketone;  
 (phenyl)-[4-(5-chloropyridin-2-yloxy)piperidin-1-ylsulphonylmethyl]-ketone;  
 (4-chlorophenyl)-(piperazin-1-ylsulphonylmethyl)-ketone;  
 (4-chlorophenyl)-[4-(*t*-butoxycarbonyl)piperazin-1-ylsulphonylmethyl]-ketone;  
 (4-hydroxyphenyl)-(morpholinosulphonylmethyl)-ketone; or  
 (phenyl)-(1,2,3,4-tetrahydroisoquinolin-2-ylsulphonylmethyl)-ketone; ~~and with the proviso that~~  
 when  $R^2$  and  $R^3$  are hydrogen,  $m$  is 0, and Ring B is 4-methylpiperazin-1-yl, then  $(R^1)_n$  is not  
 hydrogen, 4-fluoro, 4-nitro, 3,4-dimethoxy, 4-methoxy, 4-*t*-butyl, 4-trifluoromethyl, or 4-chloro;  
~~and with the proviso that~~  
 when  $R^2$  and  $R^3$  are hydrogen,  $m$  is 0, and Ring B is morpholino, then  $(R^1)_n$  is not hydrogen,  
 4-dimethylamino, 4-nitro, 4-methoxy, 4-*t*-butyl, 4-trifluoromethyl, or 4-fluoro or 4-chloro.

12. (Currently Amended) A compound of formula (Ik):



wherein:

$R^1$  is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl,  $C_{1-6}$ alkyl,  $C_{2-6}$ alkenyl,  $C_{2-6}$ alkynyl,  $C_{1-6}$ alkoxy,  $C_{1-6}$ alkanoyl,  $C_{1-6}$ alkanoyloxy,  $N$ -( $C_{1-6}$ alkyl)amino,  $N,N$ -( $C_{1-6}$ alkyl) $_2$ amino,  $C_{1-6}$ alkanoylamino,  $N$ -( $C_{1-6}$ alkyl)carbamoyl,  $N,N$ -( $C_{1-6}$ alkyl) $_2$ carbamoyl,  $C_{1-6}$ alkylS(O) $_a$  wherein  $a$  is 0 to 2,  $C_{1-6}$ alkoxycarbonyl,  $N$ -( $C_{1-6}$ alkyl)sulphamoyl,  $N,N$ -( $C_{1-6}$ alkyl) $_2$ sulphamoyl,  $C_{1-6}$ alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclyl $C_{0-6}$ alkylene-Y-, and heterocyclyl $C_{0-6}$ alkylene-Y-; or two  $R^1$  groups on adjacent carbons may form an oxy $C_{1-4}$ alkoxy group or a  $C_{3-5}$ alkylene group; wherein  $R^1$  may be optionally substituted on carbon ~~by~~ with one or more  $R^7$  groups ~~selected from  $R^7$~~ ; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by~~ with an  $R^8$  group ~~selected from  $R^8$~~ ;

$n$  is 0-3; wherein the values of  $R^1$  may be the same or different;

$R^2$  and  $R^3$  are independently selected from hydrogen, hydroxy, amino, cyano,  $C_{1-4}$ alkyl,  $C_{1-4}$ alkoxy,  $N$ -( $C_{1-4}$ alkyl)amino,  $N,N$ -( $C_{1-4}$ alkyl) $_2$ amino,  $C_{1-4}$ alkylS(O) $_a$  wherein  $a$  is 0 to 2,

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C<sub>1-4</sub>alkoxycarbonyl, C<sub>1-4</sub>alkoxycarbonylamino, C<sub>1-4</sub>alkanoyloxy, carbocyclyl, heterocyclyl, carbocyclylC<sub>1-4</sub>alkyl, and heterocyclylC<sub>1-4</sub>alkyl; or

R<sup>2</sup> and R<sup>3</sup> together form oxo or a spiro attached heterocyclyl; wherein R<sup>2</sup> and R<sup>3</sup> may be independently optionally substituted on carbon ~~by~~ with one or more R<sup>9</sup> ~~groups selected from R<sup>9</sup>~~; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by~~ with an R<sup>10</sup> ~~group selected from R<sup>10</sup>~~;

Ring B is carbocyclyl or heterocyclyl; wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by~~ with an R<sup>17</sup> ~~group selected from R<sup>17</sup>~~;

R<sup>6</sup> is a substituent on carbon and is selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy, C<sub>1-4</sub>alkyl, C<sub>2-4</sub>alkenyl, C<sub>2-4</sub>alkynyl, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkanoyloxy, N-(C<sub>1-4</sub>alkyl)amino, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkanoylamino, N-(C<sub>1-4</sub>alkyl)carbamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-4</sub>alkoxycarbonyl, N-(C<sub>1-4</sub>alkyl)sulphamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>sulphamoyl, C<sub>1-4</sub>alkylsulphonylamino, carbocyclyl, heterocyclyl, carbocyclylC<sub>0-4</sub>alkylene-Y-, and heterocyclylC<sub>0-4</sub>alkylene-Y-; wherein R<sup>6</sup> may be optionally substituted on carbon ~~by~~ with one or more R<sup>18</sup> ~~groups selected from R<sup>18</sup>~~; and wherein if said heterocyclyl contains an -NH- moiety, that nitrogen may be optionally substituted ~~by~~ with an R<sup>19</sup> ~~group selected from R<sup>19</sup>~~;

m is 0-3; wherein the values of R<sup>6</sup> may be the same or different;

Y is -S(O)<sub>a</sub>-, -O-, -NR<sup>20</sup>-, -C(O)-, -C(O)NR<sup>21</sup>-, -NR<sup>22</sup>C(O)-, or -SO<sub>2</sub>NR<sup>23</sup>-; wherein a is 0 to 2;

R<sup>7</sup>, R<sup>9</sup>, and R<sup>18</sup> are independently selected from halo, nitro, cyano, hydroxy, amino, carboxy, carbamoyl, mercapto, sulphamoyl, trifluoromethyl, trifluoromethoxy, C<sub>1-4</sub>alkyl, C<sub>2-4</sub>alkenyl, C<sub>2-4</sub>alkynyl, C<sub>1-4</sub>alkoxy, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkanoyloxy, N-(C<sub>1-4</sub>alkyl)amino, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>amino, C<sub>1-4</sub>alkanoylamino, N-(C<sub>1-4</sub>alkyl)carbamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>carbamoyl, C<sub>1-4</sub>alkylS(O)<sub>a</sub> wherein a is 0 to 2, C<sub>1-4</sub>alkoxycarbonyl, N-(C<sub>1-4</sub>alkyl)sulphamoyl, N,N-(C<sub>1-4</sub>alkyl)<sub>2</sub>sulphamoyl, C<sub>1-4</sub>alkylsulphonylamino, carbocyclyl, and heterocyclyl; wherein R<sup>7</sup>, R<sup>9</sup>, and R<sup>18</sup> may be independently optionally substituted on carbon ~~by~~ with one or more R<sup>26</sup> ~~groups~~ groups;

R<sup>8</sup>, R<sup>10</sup>, R<sup>17</sup>, and R<sup>19</sup> are independently selected from C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkanoyl, C<sub>1-4</sub>alkylsulphonyl, C<sub>1-4</sub>alkoxycarbonyl, carbamoyl, N-(C<sub>1-4</sub>alkyl)carbamoyl, N,N-(C<sub>1-4</sub>alkyl)carbamoyl, benzyl, benzyloxycarbonyl, benzoyl, carbocyclyl, heterocyclyl, and

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phenylsulphonyl; wherein  $R^8$ ,  $R^{10}$ ,  $R^{17}$ , and  $R^{19}$  may be independently optionally substituted on carbon ~~by~~ with one or more  $R^{27}$  groups;

$R^{16}$ ,  $R^{20}$ ,  $R^{21}$ ,  $R^{22}$ , and  $R^{23}$  are independently selected from hydrogen, phenyl,  $C_{1-4}$ alkylsulphonyl, and  $C_{1-4}$ alkyl;

$R^{26}$  and  $R^{27}$  are independently selected from selected from halo, nitro, cyano, hydroxy, trifluoromethoxy, trifluoromethyl, amino, carboxy, carbamoyl, mercapto, sulphamoyl, methyl, ethyl, methoxy, ethoxy, acetyl, acetoxymethyl, methylamino, ethylamino, dimethylamino, diethylamino, *N*-methyl-*N*-ethylamino, acetylamino, *N*-methylcarbamoyl, *N*-ethylcarbamoyl, *N,N*-dimethylcarbamoyl, *N,N*-diethylcarbamoyl, *N*-methyl-*N*-ethylcarbamoyl, methylthio, ethylthio, methylsulphanyl, ethylsulphanyl, mesyl, ethylsulphonyl, methoxycarbonyl, ethoxycarbonyl, *N*-methylsulphamoyl, *N*-ethylsulphamoyl, *N,N*-dimethylsulphamoyl, *N,N*-diethylsulphamoyl, and ~~or~~ *N*-methyl-*N*-ethylsulphamoyl;

or a pharmaceutically acceptable salt thereof;

with the proviso that said compound is not

(phenyl)-(5-methylpyrazol-3-ylaminosulphonylmethyl)-ketone;

(phenyl)-[(2-methyl-6-methoxy-2,3-dihydrobenzofuran-4-yl)aminosulphonylmethyl]-ketone;

(phenyl)-(1-phenyl-3-methylpyrazol-5-ylaminosulphonylmethyl)-ketone;

(phenyl)-[1-(cyclohexyl-*N*-methylaminosulphonyl)ethyl]-ketone;

(phenyl)-[1-(phenyl-*N*-methylaminosulphonyl)ethyl]-ketone;

(phenyl)-(cyclohexylaminosulphonylmethyl)-ketone;

(phenyl)-[(2-phenyl-4-acetyl-5-methylimidazol-3-yl)-*N*-methylaminosulphonylmethyl]-ketone;

(phenyl)-[(2-phenyl-4-acetyl-5-methylimidazol-3-yl)aminosulphonylmethyl]-ketone;

(phenyl)-(2,4,5,6,7,8-hexahydrocycloheptapyrazol-3-ylaminosulphonylmethyl)-ketone;

(phenyl)-(4,5,6,7-tetrahydro-2H-indazol-3-ylaminosulphonylmethyl)-ketone;

(phenyl)-[(4-phenyl-5-methylpyrazol-3-yl)aminosulphonylmethyl]-ketone;

(phenyl)-[3-(1-carboxymethyl-3-methyl-4-oxo-1,2,3,4-tetrahydrophthalazin-2-yl)anilinosulphonylmethyl]-ketone;

(phenyl)-{3-[1-(methoxycarbonylmethyl)-3-methyl-4-oxo-1,2,3,4-tetrahydrophthalazin-2-yl]anilinosulphonylmethyl}-ketone; (phenyl)-(4-methylanilinosulphonylmethyl)-ketone;

(phenyl)-(2-benzoyl-4-chloroanilinosulphonylmethyl)-ketone;

(phenyl)-(2,3-dimethylanilinosulphonylmethyl)-ketone;

(phenyl)-(3,4-dimethylanilinosulphonylmethyl)-ketone;

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(phenyl)-(3-methylanilinosulphonylmethyl)-ketone;  
(phenyl)-(3-methoxyanilinosulphonylmethyl)-ketone;  
(phenyl)-(anilinosulphonylmethyl)-ketone; (phenyl)-(2-acetylanilinosulphonylmethyl)-ketone; or  
(phenyl)-[ $\alpha$ -(N-ethylanilinosulphonyl)benzyl]-ketone.

13. (Currently Amended) A pharmaceutical composition which comprises a compound of formula (I), (II) or (III), or a pharmaceutically acceptable salt thereof, as claimed in any one of claims 9, 11 or 12, or a pharmaceutically acceptable salt thereof, in association with a pharmaceutically acceptable diluent or carrier.

14. (Currently Amended) A compound of the formula (I), (II) or (III), or a pharmaceutically acceptable salt thereof, as claimed in method for inhibiting 11 $\beta$ HSD1, comprising administering to a warm-blooded animal, a therapeutically effective amount of a compound of any one of claims 9, 11, or 12, for use in a method of prophylactic or therapeutic treatment of a warm-blooded animal, such as man.

15-16. (Cancelled).

17. (Currently Amended) A method for the treatment of a metabolic syndrome, comprising inhibiting 11 $\beta$ HSD1 ~~The use of a compound as claimed in any one of claims claim 1-8, or 10 or 16 wherein production of, or producing an, 11 $\beta$ HSD1 inhibitory effect refers to the treatment of metabolic syndrome.~~

18. (Currently Amended) A method for the treatment of a disease selected from ~~The use of a compound as claimed in any one of claims 1-8, 10 or 16 wherein production of, or producing an, 11 $\beta$ HSD1 inhibitory effect refers to the treatment of diabetes, obesity, hyperlipidaemia, hyperglycaemia, hyperinsulinemia, and or hypertension, comprising inhibiting 11 $\beta$ HSD1 as claimed in claim 1 or 10 particularly diabetes and obesity.~~

19. (Currently Amended) A method for the treatment of a disease selected from ~~The use of a compound as claimed in any one of claims 1-8, 10 or 16 wherein production of, or producing an, 11 $\beta$ HSD1 inhibitory effect refers to the treatment of glaucoma, osteoporosis, tuberculosis,~~

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dementia, cognitive disorders or depression, comprising inhibiting 11βHSD1 as claimed in claim 1 or 10.

20. (Cancelled).